(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization

International Bureau



TO THE STITLE OF BEHIND HER EXPLESIVE COLUMN CO

(43) International Publication Date 12 February 2004 (12.02.2004)

(10) International Publication Number WO 2004/013280 A3

- C12O 1/68. (51) International Patent Classification7: C12P 19/34, C12N 15/63, 15/88, C07H 21/02, 21/04, A01N 43/04
- (21) International Application Number:

PCT/US2003/016887

- 26 May 2003 (26.05.2003) (22) International Filing Date:
- (25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

5 August 2002 (05.08.2002) US 10/212,322 17 December 2002 (17.12.2002) US 10/322,086 5 May 2003 (05.05.2003) US 10/430,351

(63) Related by continuation (CON) or continuation-in-part (CIP) to earlier application:

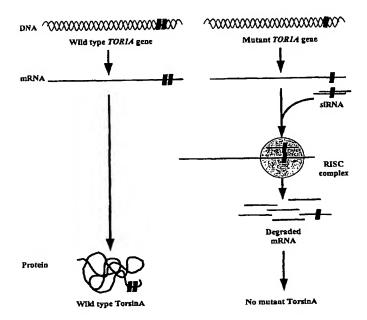
10/430,351 (CIP) US 5 May 2003 (05.05.2003) Filed on

(71) Applicant (for all designated States except US): UNIVER-SITY OF IOWA RESEARCH FOUNDATION [US/US]; 100 Oakdale Campus, 214 Technology Information Center, Iowa City, IA 52242-5000 (US).

- (72) Inventors; and
- (75) Inventors/Applicants (for US only): DAVIDSON, Beverly, L. [US/US]; 3640 Johnson Way N.E., North Liberty, IA 52242 (US). GONZALEZ-ALEGRE, Pedro [US/US]; 1930 Calvin Avenue, Iowa City, IA 52246-3100 (US). MILLER, Victor [US/US]; 1220 3rd Avenue, Iowa City, IA 52240-2011 (US). PAULSON, Henry [US/US]; 416 North Linn Street, Iowa City, IA 52245-2155 (US). HARPER, Scott [US/US]; 1 Warwick Circle, Iowa City, IA 52240-2847 (US).
- (74) Agent: VIKSNINS, Ann, S.; Fish & Richardson P.C., P.A., 60 South Sixth Street, Suite 3300, Minneapolis, MN 55402 (US).
- (81) Designated States (national): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.
- (84) Designated States (regional): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW),

[Continued on next page]

(54) Title: ALLELE-SPECIFIC siRNA-MEDIATED GENE SILENCING



(57) Abstract: The present invention is directed to small interfering RNA molecules (siRNA) targeted against an allele of interest, and methods of using these siRNA molecules.

WO 2004/013280 A3



Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

Published:

- with international search report
- before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments

(88) Date of publication of the international search report: 29 December 2005

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

INTERNATIONAL SEARCH REPORT

Interactional Application No

PCT/US03/16887

		FC1/0303/1088/			
A. CLASSIFICATION OF SUBJECT MATTER					
IPC(7) : C12Q 1/68; C12P 19/34; C12N 15/63, 15/88; C07H 21/02, 21/04; A01N 43/04					
US CL : 435/6, 91.1, 91.31, 455, 458; 536/23.1, 24.5; 514/44					
According to International Patent Classification (IPC) or to both national classification and IPC					
B. FIEL	DS SEARCHED				
Minimum documentation searched (classification system followed by classification symbols)					
U.S.: 435/6, 91.1, 91.31, 455, 458; 536/23.1, 24.5; 514/44					
	,,,,,,,		ĺ		
		 			
Documentation	on searched other than minimum documentation to the	e extent that such documents are included	in the fields searched		
1					
1					
					
	ta base consulted during the international search (nan	ne of data base and, where practicable, sea	rch terms used)		
West, Dialog, STIC sequence search					
•		·			
C. DOCUMENTS CONSIDERED TO BE RELEVANT					
Category *	Citation of document, with indication, where a	entropriate of the relevant necroses	Relevant to claim No.		
Y					
1	US 6,387,616 (ORZELIUS et al.) 14 May 2002 (ab: ID NO. 24).	suavi, vol. 1, 1, 12-14, 17, 51-52, SEQ	·1-17, 25-27		
Y	FIRE, A. et al., Potent and specific genetic interfere	nog by double stranded DNA in C	1 17 25 27		
*	elegans, Nature, Vol. 391, pages 806-811, Feb. 19,		1-17, 25-27		
	and Table 3 on p. 809).	1556, esp. at p. 600-7, fig. 1 of p. 606			
	and 10010 0 on p. 000).		ļ.		
Y	US 5,902,880 (THOMPSON) 11 May 1999 (eol. 6-	11 claims 11-16 19-24)	1-17, 25-27		
-	05 5,502,500 (1110,112,014) 11 1111, 1235 (661. 04.	11, 01111111111111111111111111111111111	1-17, 25-27		
Y	US 2002/0086356 (TUSCHL et al.) 4 July 2002 (p.	8. fig. 14h. np. 10, 27, 28)	3-8		
-	(1 00 2002) (1 00 0 1 1 0 0 0 0 0 0 0 0 0 0 0 0 0 0	o, ag. 140, pp. 10, 27, 20).	3-0		
		j			
		i			
	•		1		
		j	į		
			I		
			į.		
			i		
			j		
	1 11 11 11 11 11 11 11				
Further	documents are listed in the continuation of Box C.	See patent family annex.			
* S ₁	Special categories of cited documents: T" later document published after the international filling date or priority do				
"A" document	defining the general state of the art which is not considered to be of	and not in conflict with the application be principle or theory underlying the inventi			
particular	relevance		1		
"B" earlier app	dication or patent published on or after the international filing date	"X" document of particular relevance; the ela- considered novel or cannot be considered			
		when the document is taken alone	Lott o our miserrane such		
	which may throw doubts on priority claim(s) or which is cited to he publication date of another citation or other special reason (as	"Y" document of particular relevance: the clai	imad invention are + 6 -		
specified)	to brompount one or montel everten or emit sheets tossen (sz	"Y" document of particular relevance; the claic considered to involve an inventive step w			
	mfining to an antificial and a substitute and the s	with one or more other such documents,			
"O" document referring to an oral disclosure, use, exhibition or other means		to a person skilled in the art			
"P" document published prior to the international filing date but later than the		"&" document member of the same patent for	nily		
priority date claimed					
Date of the actual completion of the international search Date of mailing of the international search report			breport		
	-	31 UCT 2	ไ		
	2005 (14.09.2005)	- U 1 - U 1 -			
Name and mailing address of the ISA/US		Authorized officer . // Dhat	ind		
Mail Stop PCT, Attn: ISA/US Commissioner of Patents		Authorized officer Jane Zara Telephone No. (701) 208-0196			
P.O. Box 1450		0,			
	Alexandria, Virginia 22313-1450 Telephone No. (703) 208-0196				
Facsimile No. (703) 305-3230					

Form PCT/ISA/210 (second sheet) (July 1998)

INTERNATIONAL SEARCH REPORT

Interrestimal application Ma.	
PCT/US03/16887	

The YOU A STATE OF THE STATE OF
Box I Observations where certain claims were found unsearchable (Continuation of Item 1 of first sheet)
This international report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. Claim Nos.: because they relate to subject matter not required to be searched by this Authority, namely. Claims 18-24 and 41-60 were not searchable because they were multiply dependent claims.
2. Claim Nos.: 18-24 and 41-60 because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically.
Claim Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II Observations where unity of invention is lacking (Continuation of Item 2 of first sheet)
This International Searching Authority found multiple inventions in this international application, as follows: Please See Continuation Sheet 1.
4. No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.: 1-17,25-27 and 61-81 Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.

Form PCT/ISA/210 (continuation of first sheet(1)) (July 1998)